The effects of pneumoperitoneum pressure on blood gases, respiratory and venous systems during laparoscopic cholecystectomy: A prospective randomized trial

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ABSTRACT

Introduction: Increased abdominal pressure during pneumoperitoneum may distress respiratory functions and venous systems. The aim of this study was to evaluate the effect of low and high pneumoperitoneum pressure during laparoscopic cholecystectomy.

Materials and Methods: Total of 40 patients were randomized for use of either low (8 mmHg) or high (14 mmHg) pneumoperitoneum pressure. Respiratory mechanics were monitored continuously, arterial blood gases were analyzed via radial artery catheter, and duplex scan of left common femoral vein was performed. Ten days after surgery, venous duplex scan of lower limbs was used to detect signs of deep vein thrombosis.

Results: While peak inspiratory pressure significantly increased with low and high pneumoperitoneum pressure, dynamic compliance significantly decreased. Although carbon dioxide insufflation caused decrease in blood pH in both groups, it was only significant at high pneumoperitoneum pressure. Duplex scan of femoral vein revealed significant increase in diameter and decrease in peak blood velocity at high pneumoperitoneum pressure.

Conclusion: Respiratory acidosis may occur due to decreased compliance, and pneumoperitoneum causes reversible venous stasis, especially during use of high pressure. Results indicated that performing laparoscopy with lower pneumoperitoneum pressure decreased these adverse effects, especially in patients with cardiopulmonary comorbid diseases. Prophylaxis for venous thromboembolism in high-risk patients undergoing laparoscopic cholecystectomy is recommended.

Keywords: Acid-base balance; deep vein thrombosis; laparoscopy; pneumoperitoneum; pulmonary mechanics; venous stasis.
Introduction

Laparoscopy with carbon dioxide (CO₂) pneumoperitoneum significantly influences blood gases, respiratory and deep venous systems. Although there are many experimental studies performed on animal models, the influence of different pneumoperitoneum pressures on patients’ blood gases, respiratory, and venous systems is not clear and some effects of pneumoperitoneum on human body still remains controversial.[1–10] Intraoperative venous stasis may increase the risk for preoperative or postoperative deep vein thrombosis and pulmonary embolism, especially during laparoscopic procedures.[8,11–15] Some authors favor venous thromboembolism (VTE) prophylaxis, whereas some not.[12,13,16] This paper reports the result of a prospective study investigating the effect of different pneumoperitoneum pressures (8 mmHg or 14 mmHg) on arterial blood gas, respiratory parameters, and venous system during laparoscopic cholecystectomy (LC).

Materials and Methods

All patients evaluated for the laparoscopic treatment of acute cholecystitis were considered to take part in the trial. Patients over 60 years of age with pulmonary or cardiovascular disease, previous surgeries, a history of venous or arterial diseases were excluded. All patients were in class I and II according to the American Society of Anesthesiology (ASA) classification.

The local ethics committee approved the study protocol. Following written valid consents of the patients, they were randomly assigned to LC under 8 mmHg (Group I) or 14 mmHg (Group II) pneumoperitoneum pressures (PP). Randomization was performed using sealed envelopes. A total of 40 patients were randomized as 20 being in each group. Hospital stay, operation time, mortality and morbidity rate were recorded.

CO₂ pneumoperitoneum was executed by a Veress needle initially at a slow flow (1 L/min) and then faster flow (maximum 20 L/min) to avoid a vasovagal reaction. Before insufflation, during pneumoperitoneum and following desufflation, respiratory mechanics were recorded and dynamic compliance, peak inspiratory pressure, and arterial blood gases were analyzed by a blood gas analyzer from the radial artery catheter. Arterial blood samples were taken puncturing the left radial artery prior to pneumoperitoneum, 20 minutes after pneumoperitoneum and 20 minutes after desufflation. The following parameters of acid-base balance were recorded: pH, PaCO₂, PaO₂, HCO₃⁻ base excess.

Peak blood velocity and diameter of left common femoral vein were measured by color duplex scanning before insufflation, during pneumoperitoneum and following desufflation. Ten days after surgery, lower limbs were examined by a venous duplex scanning to detect signs of deep vein thrombosis.

Anesthesia was induced with intravenous (IV) propofol (2 mg/kg) and fentanyl (1 μg/kg), followed by vecuronium (0.1 mg/kg) for intubation and carried with sevoflurane (1 MAC minimum alveolar concentration).

A ventilator was used for artificial lung ventilation at a rate of 12 breaths/min and tidal volume of 10 mL/kg, with a mixture of air and oxygen (FiO₂ 40%). All patients received prophylactic antibiotic as cefazolin sodium 1 g. The fourtrocar technique was preferred and all patients were operated on in 15° reverse Trendelenburg position with slightly (10°) rotated to the left side.

Instat Statistical Package (GraphPad, CA, USA) was used for statistical analysis. Suitable data were expressed as mean ± standard deviations. Student’s t-test and ANOVA were used for statistical analysis. Statistical significance level was established as p<0.05.

Results

The groups were well matched and there were no significant differences of age, sex, body mass index (BMI), operating time, anesthesia time, ASA class, and mortality and morbidity rate between the groups (Table 1). Although mean operating time and anesthesia time were longer in Group I (8 mmHg-PP) than in Group II (14 mmHg-PP), they were not statistically significant.

There were no serious complications in both groups. Wound infection was observed in one patient in Group I and one in Group II had fever secondary to atelectasis.

There was no mortality in both groups. The effects of PP with low and high pressures on arterial blood gases, acid-base balance (ABB), respiratory and deep venous systems are compared at Table 2.

Arterial Blood Gases and ABB

Blood pH decreased slightly by 8 and 14 mmHg after intraperitoneal insufflation which was also noticed after desufflation. Only in Group II, the decrease between PP and before insufflation was statistically significant (p<0.001). However, a gradual decrease toward normal values (pH
7.35–7.45) was noticed. The PaCO₂ levels considerably increased with pneumoperitoneum. While this wasn’t significant during PP in both groups, it was significant after desufflation. In the measurement of arterial PaCO₂ fit pH values, no significant differences were found between the groups by a mean of PaO₂ at the same steps of operation. No statistically significant differences were found between the groups and comparisons in the groups by mean of PaO₂.

In Group I, there was no significant difference among pre-insufflation, during insufflation and after desufflation by mean of HCO₃⁻ (p>0.05). However, in Group II, bicarbonate level decreased moderately during insufflation and continued till after insufflation. In this group, there was a significant difference between pre-insufflation and during insufflation (p<0.001), during insufflation and after desufflation (p<0.05). There was no significant decrease during insufflation in two groups by mean of HCO₃⁻; however, it was noticed after desufflation in Group II rather than in Group I. All BE values were found in normal range. Statistical difference was recorded in Group II during insufflation and after desufflation. Although BE was lower in Group II than in Group I, no significant differences were found between the two groups by mean of BE.

### Respiratory Parameters

Peak inspiratory pressure (PIP) was followed during the study. In Group I, PIP increased significantly during insufflation rather than before insufflation. After desufflation, PIP was close to the PIP values before insufflation. In addition, during insufflation, PIP increased to higher values compared to pre-insufflation which was significant in Group II (p<0.01). These changes were reversible after desufflation. No significant difference was found in two

### Table 1. Patients’ characteristics and operative data

<table>
<thead>
<tr>
<th></th>
<th>Group 1</th>
<th>Group 2</th>
<th>p</th>
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</thead>
<tbody>
<tr>
<td>Mean age (years)</td>
<td>46.3 (±8.2)</td>
<td>46.5 (±8.2)</td>
<td>&gt;0.05</td>
</tr>
<tr>
<td>Sex (male/female)</td>
<td>6/14</td>
<td>5/15</td>
<td>&gt;0.05</td>
</tr>
<tr>
<td>BMI</td>
<td>28.7 (22.4–36.3)</td>
<td>27.1 (23.2–35.4)</td>
<td>&gt;0.05</td>
</tr>
<tr>
<td>ASA class (I/II)</td>
<td>13/7</td>
<td>14/6</td>
<td>&gt;0.05</td>
</tr>
<tr>
<td>Operating time (minutes)</td>
<td>62.8±25.6</td>
<td>55.7±18.4</td>
<td>&gt;0.05</td>
</tr>
<tr>
<td>Anesthesia time (minutes)</td>
<td>72.8±23.2</td>
<td>56.9±16.9</td>
<td>&gt;0.05</td>
</tr>
<tr>
<td>Morbidity rate (%)</td>
<td>5</td>
<td>5</td>
<td>&gt;0.05</td>
</tr>
</tbody>
</table>

Values are mean±standard deviation.

### Table 2. The effects of different insufflation pressures on arterial blood gases, acid-base balance (ABB), respiratory and deep venous systems

<table>
<thead>
<tr>
<th></th>
<th>Before insufflation</th>
<th>During pneumoperitoneum</th>
<th>After desufflation</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td>Group 1</td>
<td>Group 2</td>
<td>Group 1</td>
</tr>
<tr>
<td>DC (mL/cmH₂O)</td>
<td>30±4.9</td>
<td>32.1±6.3**</td>
<td>26.6±5.4**</td>
</tr>
<tr>
<td>PIP (mmHg)</td>
<td>24.2±4.4</td>
<td>24.2±3.67**</td>
<td>27.1±5.17**</td>
</tr>
<tr>
<td>pH</td>
<td>7.41±0.04</td>
<td>7.44±0.04</td>
<td>7.39±0.03</td>
</tr>
<tr>
<td>PaCO₂</td>
<td>33.35±3.31</td>
<td>31.54±3.47</td>
<td>34.99±3.86</td>
</tr>
<tr>
<td>PaO₂</td>
<td>252.60±33.52</td>
<td>266.80±67.30</td>
<td>247.90±48.79</td>
</tr>
<tr>
<td>HCO₃⁻</td>
<td>24.47±2.97</td>
<td>24.10±0.96</td>
<td>24.58±5.04</td>
</tr>
<tr>
<td>BE</td>
<td>0.12±1.46</td>
<td>-0.73±1.48</td>
<td>0.09±1.73</td>
</tr>
<tr>
<td>FV flow speed (cm/sc)</td>
<td>17.2±8.1</td>
<td>24.6±3.8**</td>
<td>16.6±8.2</td>
</tr>
<tr>
<td>FV diameter (mm)</td>
<td>11.04±3.1</td>
<td>12.5±1.31</td>
<td>11.14±3.21</td>
</tr>
</tbody>
</table>

*p<0.05 between groups. **p<0.05 in group.
groups by mean of PIP, recorded at the same steps of the surgery. Dynamic compliance (TV/PlatoP-PEEP) was decreased during insufflation, compared to before insufflation in both groups. During insufflation (p<0.05) and after desufflation dynamic compliance decreased to normal values at the beginning in Group I. Same changes were detected in Group II. The significant decrease in DC was determined during insufflation, rather than before insufflation (p<0.05). In this group, after desufflation, dynamic compliance decreased to values prior to insufflation. DC changes at the same steps of the operation were found insignificant.

Venous System

Left femoral vein diameter was consistent with peak blood velocity. In Group I, an increase on left femoral vein diameter and a decrease in peak blood velocity per second during insufflation were observed; however, these changes were not statistically significant. In group II, although an increase in left femoral vein diameter was statistically insignificant, a significant decrease in peak velocity per second was revealed (p<0.001). In both groups, after desufflation, femoral vein diameter returned to the values prior to insufflation. An increase in the left femoral vein diameter and a decrease in peak blood velocity per second during insufflation in Group II was significantly different from Group I; however, no statistically significant changes were determined following desufflation in both groups by mean of diameter and peak blood velocity. Ten days after surgery, lower limb venous duplex scanning and one deep vein thrombosis were detected in each group. LMWH was used for the treatment of these two patients.

Discussion

Currently, laparoscopic surgery has gained widespread acceptance for the diagnosis and treatment of many diseases. Insufflation is, one of the basic features of laparoscopic surgery, provides the space needed for best view and optimal working. Since laparoscopic surgery needs smaller incisions and less dissections when compared to open surgery, patients have less postoperative pain and less complications including wound infection, respiratory and gastrointestinal complications. Additionally, it is preferred by surgeons and patients as it provides better cosmetic outcomes reducing hospitalization time and shortening the time needed to return to daily activities.

However, PP with CO₂ insufflation, the basic process of laparoscopic surgery, is accompanied by a certain number of pathophysiological changes which are suggested to be in relation with increases in intraabdominal pressure (IAP) and CO₂ absorption through peritoneum and its transition into the systemic circulation. Although there are many experimental and clinical studies regarding these effects of PP, its influences on arterial blood gases, respiratory and deep venous systems are unclear and its effects on human body are still controversial.[1,10]

Increased IAP is associated with diaphragm elevation causing an increase in the intrathoracic pressure while resulting in an increase in the diaphragm tension and restraining lung expansion. Therefore, compression atelectasia occurs. As the number of ventilated alveoli decrease, dead space expands and causes the lung functional residual capacity to decrease. As a result, ventilation-perfusion mismatch appears. As IAP increases, dynamic compliance reduces whereas airway pressure, PP, and plateau pressures increase.[17–22] In our study, significant decreases in DC and increases in PIP during PP in both high and low PP groups were detected. In reference to the high PP group, decrease in DC and increase in PIP were less significant in low PP group. Following desufflation, DC and PIP values returned to initial values before insufflation. Proportionately, with the increase in IAP, significant changes in DC and PIP were detected.

CO₂, the ideal gas to compose PP, reaches the maximum levels in circulation within a short time after being absorbed rapidly through peritoneum. Some researchers suggest that CO₂ absorption time increases proportionately with the operation time and IAP increase; however, others report that there is no proportional relationship between IAP and peritoneal absorption of CO₂[18,23] CO₂ elimination occurs directly from lungs by ventilation. Elimination is directly proportional with cardiac output and ventilation rate. Supporters of the idea that increases in IAP and CO₂ absorption are directly proportional suggest that CO₂ elimination through lungs is limited as a result of increased intrathoracic pressure, decreased cardiac output, limitation of the expansion capacity of lungs, occurrence of compression atelectasia, and ventilation-perfusion mismatch due to diaphragm elevation.

The researchers supporting the idea that trans-peritoneal absorption of CO₂ reduces as the IAP increases suggest that IAP limits absorption of CO₂ through capillary vessels with compression impact on peritoneal capillary vessels. In our study, despite not being statistically significant,
PaCO₂ increased during PP and more during high pressure insufflations. Following desufflation, PaCO₂ increased in both groups, more in high insufflation pressures, as well. Although mean operation and anesthesia time were shorter, PaCO₂ values increased more significantly during high insufflation pressures in comparison with low pressures indicating the linear association between IAP and PaCO₂.

Elimination of CO₂ from circulation occurs directly by ventilation through lungs. Insufficient ventilation leads to CO₂ aggregation, especially in bones and muscles. Significant increase in PaCO₂ following desufflation may be a result of CO₂ release from tissues into circulation. Insufficient ventilation may cause hypercapnia and acidosis. Persistent hypercapnia induces renal response including H+ secretion from renal tubuli and bicarbonate passage into the extracellular zone. As almost all CO₂ is eliminated during laparoscopy, compensatory hyperventilation is required in order to prevent hypercapnia and acidosis.

Intra-abdominal pressure changes during insufflation have influences on ABB. This may cause major problems during and after surgery especially in patients with respiratory and cardiac co-morbidities. High IAP may result in acid-base imbalance increasing CO₂ absorption through peritoneum and limiting CO₂ elimination by ventilation particularly with an increase in operation time. Some experimental and clinical studies suggest that abdominal pressure formed by CO₂ changes ABB towards acidosis and hypercapnia. Although the mechanisms causing these changes are still unclear, predominant opinions suggest acidosis associated with trans-peritoneal CO₂ absorption rather than negative effects of increased IAP on ventilation.

Sefr and colleagues indicate that there is no statistically significant difference in ABB between 10 mmHg and 15 mmHg pressures in a clinical study on ASA I and ASA II patients. In our study, a slight decrease in blood pH was detected following both low and high pressure pneumoperitoneum, which was also observed after desufflation.

Statistically significant decrease was detected only in high pressure pneumoperitoneum group. Despite this decrease, pH ranged between normal values (7.35–7.45). In high pressure group, statistically significant (p<0.001) decrease in HCO₃ parallel to changes in pH was identified, which continued significantly after desufflation (p>0.05). In low pressure group, statistically insignificant decrease in HCO₃ during pneumoperitoneum and after desufflation was detected. Although all BE measurements ranged between normal values, BE values during insufflation and after desufflation showed significant changes only in high pressure group. These results indicate negative effects of increased IAP on respiratory system besides arterial blood gases and ABB. Although these pathophysiological changes do not cause abnormalities in patients with normal respiratory functions, they may result in hypoxia in patients with COPD or emphysema.

However, in laparoscopic interventions with high IAP some cardio-vascular changes also occur and cause serious problems in patients with low cardiac reserve. These changes are infrequent in interventions with low IAP. We suggest that high risk patients with cardiac and respiratory diseases can be operated with lower morbidity and mortality rates by obtaining well cardio-pulmonary monitoring and avoiding unnecessary patient positions.

Increased IAP leads to decrease in venous return in lower extremity veins as a result of direct compression on IVC and iliac veins. In surgical procedures practiced on anti-Trendelenburg position, as a result of gravity, compression of visceral organs on iliac veins cause deceleration in femoral blood flow. Deceleration effects of increased ICP and anti-Trendelenburg position on femoral vein flow has already been shown. It was suggested that these effects on deep venous system induce the factors composing the Virchow triad. However, some authorities do not recommend routine VTE prophylaxis.

In our study, the effects of low and high pressure PP on femoral vein blood flow velocity and diameter were compared and a statistically insignificant increase in the left common femoral vein diameter and decrease in peak blood flow velocity per second during PP were detected. With high pressure PP, statistically insignificant increase in CFV diameter and a significant (p<0.001) decrease in peak blood flow velocity per second during PP were determined. These results indicate that increased IAP causes deceleration in the blood flow of the deep venous system and stasis, the major etiological factor for DVT. Deep venous system Doppler examination of the lower extremities on the postoperative tenth day showed asymptomatic DVT in one patient in each group despite early mobilization.

Although laparoscopic surgery has currently gained widespread acceptance, debates on routine VTE prophylaxis still continue. Since the real incidence of DVT and PE af-
ter laparoscopic surgery is unknown and the number of our subjects was insufficient, it is not possible to identify the need for routine VTE prophylaxis in laparoscopic surgery accurately. However, in the light of the results of the study, routine VTE prophylaxis in patients who are at high risk for laparoscopic surgery is recommended.

**Conclusion**

Increase in intra-abdominal pressure during pneumoperitoneum affects respiratory functions, decreases dynamic compliance, and increases PIP. It changes arterial blood gases and acid-base balance towards hypercapnia and acidosis. It causes stasis in the deep venous system and correspondingly increases the risk for VTE. Patients at high risk for cardiac and respiratory diseases can be operated with low pneumoperitoneum pressures with lower morbidity and mortality. Prophylaxis for venous thromboembolism in patients carrying risk for laparoscopic cholecystectomy is recommended.

**Disclosures**

**Ethics Committee Approval:** The study was approved by the Local Ethics Committee.

**Peer-review:** Externally peer-reviewed.

**Conflict of Interest:** None declared.

**References**


